Supplementary Figure 1

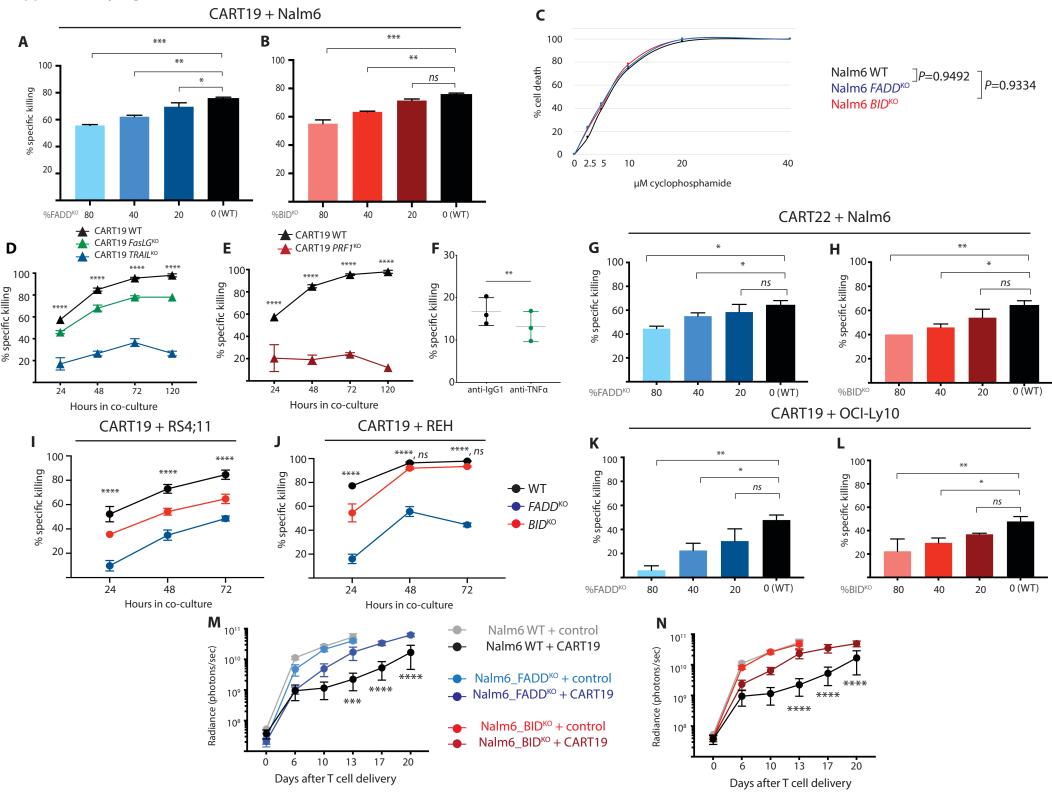


Figure S1. Impairment of death receptor signaling results in resistance to CART19.

- (A) Nalm6 FADDKO and (B) BIDKO cells were combined with Nalm6 WT cells at progressive KO concentrations, co-cultured with CART19 and Nalm6 survival was measured after 24 hours.
- (C) WT, FADDKO and BIDKO Nalm6 cells were exposed to increasing concentrations of cyclophosphamide and cell survival was measured after 24 hours.
- (D) Survival of WT Nalm6 cells cultured with CART19 cells edited for loss of death receptor ligands. Significance reflects difference between WT and TRAILKO samples.
- (E) Survival of WT Nalm6 cells cultured with perforin-deficient CART19.
- (F) Survival of WT Nalm6 cells cultured with CART19 cells in the presence of control (anti-IgG1, 1ug/mL) or anti-TNFa (1ug/mL) inhibitory antibodies. E:T ratio of 0.03 CAR T cells to 1 Nalm6.
- (G) Nalm6 FADDKO and (H) BIDKO cells were combined with Nalm6 WT cells at progressive KO concentrations, co-cultured with CART22 and Nalm6 survival was measured after 24 hours.
- (I-J) Survival over time of WT, FADDKO and BIDKO (I) RS4;11 and (J) REH cells combined with CART19.
- (K) OCI-Ly10 FADDKO and (L) BIDKO cells were combined with OCI-Ly10 WT cells at progressive KO concentrations, co-cultured with CART19 and OCI-Ly10 survival was measured after 24 hours.
- (M-N) Disease progression over time in immunodeficient mice engrafted with luciferase-expressing Nalm6 (WT, BIDKO or FADDKO) after treatment with 5x105 control or CART19 cells.
- *P<0.05, **P<0.01, ***P<0.001, ****P<0.0001, ns: not significant